

AMENDMENT TO THE SPECIFICATION

On page 1 of the specification please amend the title:

3-AMINO-~~PIPERADINE~~ PIPERIDINE DERIVATIVES AND METHODS OF  
MANUFACTURE

On page one, paragraph two:

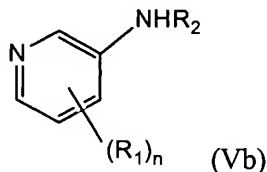
This invention relates to 3-amino-~~piperadine~~ piperidine derivatives, their intermediates, and methods of manufacture.

On page one, paragraph three:

Pyrrolo[2,3-d]pyrimidine compounds are inhibitors of protein kinases, such as the enzyme Janus Kinase 3 (JAK3) and are therefore useful therapy as immunosuppressive agents for organ transplants, xeno ~~transplation~~ transplantation, lupus, multiple sclerosis, rheumatoid arthritis, psoriasis, Type I diabetes and complications from diabetes, cancer, asthma, atopic dermatitis, autoimmune thyroid disorders, ulcerative colitis, Crohn's disease, Alzheimer's disease, Leukemia and other indications where immunosuppression would be desirable. The pyrrolo[2,3-d]pyrimidine compounds, pharmaceutical compositions thereof and methods of use are described in co-pending application serial no. 09/732669, filed December 8, 2000, and assigned to the assignee of the present invention, which is incorporated herein by reference for all purposes.

On page 6, paragraph two:

The present invention also relates to formation of the compound of the formula (IVb) by reacting a compound having the formula (Vb)



with  $(R_{14}-O-(C=O))_2O$   $(R_{14}-O-(C=O))_2O$  or  $R_{14}-O-(C=O)-X$  wherein X is halo.

On page 11, paragraph two:

In step 2 of Scheme 1, the carbamate of formula (Va) is oxidized to form the oxidation product (IVa). Typically, the oxidation reaction produces a mixture of compounds having the formula (IVa). Any suitable oxidation conditions may be used. Preferred conditions include electrochemical oxidation, such as performing the oxidation reaction in an electrolytic solution in an electric cell and electrolyzing the cell. In one embodiment, the electrolytic solution is a mixture of acetic acid and potassium acetate. In another embodiment, the electrolytic solution includes acetic anhydride. The cathode and anode may be made of any suitable material, including platinum and ~~niobium~~ niobium. The mixture is then electrolyzed at an appropriate current until

On page 24, paragraph two:

To a three neck round bottom flask was added 360g of 4-methylpiperidine, 470mL of triethylamine, and 390mL of methylene chloride and the mixture was cooled in an ice bath. To this mixture was added methylchloroformate (260mL) in methylene chloride (215mL) slowly to maintain a reaction temperature of 20C or below. The reaction was stirred overnight, then 200mL of water was added and the layers were separated. The organic layer was washed with dilute HCl, satd. ~~NaHCO<sub>3</sub>~~ NaHCO<sub>3</sub>, and brine, and then the organic layer was dried over sodium sulfate and the solvent was removed in vacuo. The product was distilled at 90-93C at ~10mm pressure to provide 338g of product.